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## Heart Failure

### EXERCISE TRAINING UP-REGULATES PHYSIOLOGICAL GROWTH SIGNALLING IN THE MYOCARDIUM OF PATIENTS WITH MECHANICAL CIRCULATORY SUPPORT: RELEVANCE TO FUNCTIONAL RECOVERY

Oral Contributions

South, Room 102

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**Background:** To investigate whether exercise training in patients with end-stage heart failure (HF) supported by Ventricular Assist Device (VAD) results to benefits in functional capacity and favorable changes in growth signalling pathways implicated in cardiac remodeling.

**Methods:** Thirty-two patients with VAD bridged to heart transplantation (HTx) were included in the study. Patients were randomly assigned to a non-training control group (CG, n=16) or to a training group (GT, n=16), aerobic training for 45 min at Borg scale 12-14 for 3-4 times a week plus high-intensity inspiratory muscle training, initially at the hospital (12 weeks) and then at the home with confirmation of adherence until HTx. Echocardiography and cardiopulmonary exercise testing (peak V<sub>O2</sub>) were performed pre VAD and pre-HTx in as well as thyroid hormone level measurements.

Total (t) and phosphorylated (p) Akt, ERK1/2 and JNK as well as TR $\alpha$ 1 protein, which is downregulated in HF, were measured in myocardial tissue by western blotting analysis, at the time of VAD placement and at HTx, in 7 pts in each group.

**Results:** Physical training resulted in a significant increase in both peak V<sub>O2</sub> (ml/kg/min) and LVEF (%) in GT as compared to CG [17.0(sem, 0.8) vs 12.6 (0.7), p<0.05 and [29.5(1.5) vs 22.6(0.7), p<0.05] respectively. In pre-HTx samples, the ratio of p/t-AKT was 2 fold higher in GT than in group CG, p<0.05. The ratio of p/t-ERKs was 1.6 fold higher (p<0.05) and the ratio of p/t-JNK levels were 2.5 fold lower (p<0.05) in GT vs CG. A 2.0 fold increase in myocardial TR $\alpha$ 1 expression and an increase in circulating T3 was found in GT vs CG, p<0.05.

**Conclusions:** Physical training in VAD recipients improves aerobic capacity and this response is associated with favorable changes in physiologic growth signaling pathways, and the TR $\alpha$ 1/T3 concomitant increase.